### P NT COOPERATION TREAT

From the INTERNATIONAL BUREAU

PCT	To:	
NOTIFICATION OF ELECTION (PCT Rule 61.2)	Assistant Commissioner for Patents United States Patent and Trademark Office Box PCT Washington, D.C.20231 ETATS-UNIS D'AMERIQUE	
Date of mailing (day/month/year) 14 September 2000 (14.09.00)	in its capacity as elected Office	
International application No. PCT/US99/30503	Applicant's or agent's file reference B0192/7011WO	
International filing date (day/month/year) 21 December 1999 (21.12.99)	Priority date (day/month/year) 22 December 1998 (22.12.98)	
Applicant GORDON, Robert, Douglas et al		
1. The designated Office is hereby notified of its election made:    X   in the demand filed with the International Preliminary Examining Authority on:   24 May 2000 (24.05.00)   in a notice effecting later election filed with the International Bureau on:   2. The election   X   was   was not   was not   was not   made before the expiration of 19 months from the priority date or, where Rule 32 applies, within the time limit under Rule 32.2(b).		
	Authorized officer	

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland

Manu Berrod

Telephone No.: (41-22) 338.83.38

From the

INTERNATIONAL PRELIMINARY EXAMINING AUTHORITY

BALDOCK, Sharon Claire BOULT WADE TENNANT Verulam Gardens 70 Grav's Inn Road London WC1X 8BT GRANDE BRETAGNE



PCT

NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL PRELIMINARY EXAMINATION REPORT (PCT Rule 71.1)

Date of mailing (day/month/year) 19.04.2001

Applicant's or agent's file reference SCB/51935/004

IMPORTANT NOTIFICATION

International application No. PCT/US99/30503

International filing date (day/month/year) 21/12/1999

Priority date (day/month/year) 22/12/1998

Applicant

JANSSEN PHARMACEUTICA N.V. et al.

- 1. The applicant is hereby notified that this International Preliminary Examining Authority transmits herewith the international preliminary examination report and its annexes, if any, established on the international application.
- 2. A copy of the report and its annexes, if any, is being transmitted to the International Bureau for communication to all the elected Offices.
- 3. Where required by any of the elected Offices, the International Bureau will prepare an English translation of the report (but not of any annexes) and will transmit such translation to those Offices.

#### 4. REMINDER

The applicant must enter the national phase before each elected Office by performing certain acts (filing translations and paying national fees) within 30 months from the priority date (or later in some Offices) (Article 39(1)) (see also the reminder sent by the International Bureau with Form PCT/IB/301).

Where a translation of the international application must be furnished to an elected Office, that translation must contain a translation of any annexes to the international prefiminary examination report. It is the applicant's responsibility to prepare and furnish such translation directly to each elected Office concerned.

For further details on the applicable time limits and requirements of the elected Offices, see Volume if of the PCT Applicant's Guide.

RECEIVED

2 3 APR 2001

Name and mailing address of the IPEA/

European Patent Office D-80298 Munich

Tel. +49 89 2399 - 0 Tx; 523656 epmu d Fax: +49 89 2399 - 4465

Authorized officer

Emslander, S

Tel.+49 89 2399-8718

BOULT WADE TENNAME

Form PCT/IPEA/416 (July 1992)

Applicant's or agent's file reference

SCB/51935/004

PCT/US99/30503

C12N15/12

International application No.

See Notification of Transmittal of International

22/12/1998

Preliminary Examination Report (Form PCT/IPEA/416)

Priority date (day/month/year)

### PATENT COOPERATION TREATY

## **PCT**

### INTERNATIONAL PRELIMINARY EXAMINATION REPORT

FOR FURTHER ACTION

21/12/1999

International Patent Classification (IPC) or national classification and IPC

JANSSEN PHARMACEUTICA N.V. et al.

Form PCT/IPEA/409 (cover sheet) (January 1994)

International filing date (day/month/year)

1. This international preliminary examination report has been prepared by this international Preliminary Examining Authority

(PCT Article 36 and Rule 70)

and is	trans	smitted to the applicant according to Article 56.		
2. This F	EPC	PRT consists of a total of 8 sheets, including the	s cover sheet.	
h	an s	port is also accompanied by ANNEXES, i.e. sh mended and are the basis for this report and/or ule 70.16 and Section 607 of the Administrative	sets of the description, claims and/or drawings which hav sheets containing rectifications made before this Authority Instructions under the PCT).	
These	ann	exes consist of a total of sheets.		
3. This r	eport	contains indications relating to the following Iter	TIS:	
1	⊠	Basis of the report		
II	⊠	Priority		
111	⊠	Non-establishment of opinion with regard to no	ovelty, Inventive step and industrial applicability	
IV.				
<ul> <li>Heasoned statement under Article 35(2) with citations and explanations suporting such st</li> </ul>			regard to novelty, inventive step or industrial applicability; tement	
VI	$\boxtimes$	Certain documents cited		
VII	×	Certain defects in the international application		
VIII	Ø	Certain observations on the international appli	cation	
Dale of sub	missio	on of the demand	Date of completion of this report	
24/05/2000			19.04.2001	
		g address of the international ining authority:	Authorized officer	
European Patent Office D-80298 Munich Tel, 149 89 2399 - 0 Tx: 523656 epmu d			Hecki, K	
	Fax	+49 89 2399 - 4465	Telephone No. +49 89 2399 8430	

International application No. PCT/US99/30503

		is of the report				
1.	<ol> <li>With regard to the elements of the international application (Replacement sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "reginally filed" and are not annexed to this report since they do not contain amendments (Rules 70.16 and 70.17));</li> <li>Description, pages:</li> </ol>					
	1-55	5	as originally filed			
	Clai	ms, No.:				
	1-72	2	as originally filed			
	Dra	wings, sheets:	•			
	1-54	•	as originally filed			
	Sequence listing part of the description, pages:					
	1-12, as originally filed					
2.	<ol><li>With regard to the language, all the elements marked above were available or furnished to this Authority in the language in which the international application was filed, unless otherwise indicated under this item.</li></ol>					
	These elements were available or furnished to this Authority in the following language: , which is:					
	☐ the language of a translation furnished for the purposes of the international search (under Rule 23.1(b))					
			ublication of the international application (under Rule 48.3(b)).			
	the language of a translation furnished for the purposes of international preliminary examination (under Re 55.2 and/or 55.3).					
3.	<ol><li>With regard to any nucleotide and/or amino acid sequence disclosed in the international application, the international preliminary examination was carried out on the basis of the sequence listing:</li></ol>					
	Ø	contained in the in	nternational application in written form.			
	×	filed together with	the international application in computer readable form.			
		furnished subseq	uently to this Authority in written form.			
		furnished subseq	uently to this Authority in computer readable form.			
			at the subsequently furnished written sequence listing does not go beyond the disclosure In application as filed has been furnished.			
		The statement the listing has been for	at the information recorded in computer readable form is identical to the written sequence unished.			
4	The	amendments hav	e resulted in the cancellation of:			

International application No. PCT/US99/30503

			•			
		the description,	pages:			
		the claims,	Nos.:			
		the drawings,	sheets:			
5.		This report has been established as if (some of) the amendments had not been made, since they have been considered to go beyond the disclosure as filed (Rule 70.2(c)):				
		(Any replacement sheet containing such amendments must be referred to under Item 1 and annexed to this report.)				
6.	6. Additional observations, if necessary:					
H.	Pric	ority				
1.		This report has been prescribed time limit	established as if no priority had been claimed due to the failure to turnish within the the requested: $_{}$			
		☐ copy of the earl	er application whose priority has been claimed.			
		☐ translation of the	e earlier application whose priority has been claimed.			
2.		This report has been been found invalid.	established as if no priority had been claimed due to the fact that the priority claim has			
	Thus for the purposes of this report, the international filing date indicated above is considered to be the relevant date.					
3.	Additional observations, if necessary:     see separate sheet					
111	. No	ı-establishment of o	pinion with regard to novelty, inventive step and industrial applicability			
1.	<ol> <li>The questions whether the claimed invention appears to be novel, to involve an inventive step (to be non- obvious), or to be industrially applicable have not been examined in respect of:</li> </ol>					
		the entire internation	al application.			
	Ø	claims Nos. 28,29,32	2-38,51,52,61,62.			
because:						
	×		l application, or the said claims Nos. 32-38,52,61,62 relate to the following subject matter ire an international preliminary examination (specify):			
			ns or drawings (indicate particular elements below) or said claims Nos. are so unclear pinion could be formed (specify):			

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		the claims, or said claims Nos. are so inadequately supported by the description that no meaningful opinion could be formed.				
	☑ no international search report has been established for the said claims Nos. 28,29,61.					
2.	and	meaningful international preliminary examination cannot be carried out due to the tailure of the nucleotide nd/or amino acid sequence listing to comply with the standard provided for in Annex C of the Administrative structions:				
		the written form has not been furnished or does not comply with the standard.				
	the computer readable form has not been furnished or does not comply with the standard.					
٧.		leasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; Itations and explanations supporting such statement				
1.	Stat	atement				
	Nov	elty (N)	Yes: No:	Claims Claims	1-27,30-50,52-72	
	Inve	entive step (IS)	Yes: No:	Claims Claims	1-27,30-50,52-72	
	indu	strial applicability (IA)	Yes: No:	Claims Claims	1-27,30,31,39-50,63-60,63-72	
2.	Cita	tions and explanations				

## see separate sheet

Certain documents cited
 Certain published documents (Rule 70.10)

and / or

2. Non-written disclosures (Rule 70.9)

see separate sheet

### VII. Certain defects in the international application

The following defects in the form or contents of the international application have been noted: see separate sheet

### VIII. Certain observations on the international application

The following observations on the clairty of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made:

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see separate sheet

# INTERNATIONAL PRELIMINARY International application No. PCT/US99/30503 EXAMINATION REPORT - SEPARATE SHEET

### Re Item II

Priority

Documents D1-D3 were published after the priority date of the present application. They do not, therefore, constitute part of the state of the art in the meaning of Rule 64(1)(b) PCT. Since the present application seems entitled to the priority date (22.12.98) claimed these documents will also not become relevant with respect to novelty and inventiveness in any regional phase.

However, since these documents claim priority dates earlier than that of the present application, they might become relevant unde Art.54(3) and (4) EPC (see also Re Item VI).

#### Re Item III

Non-establishment of opinion with regard to novelty, inventive step and industrial applicability

- No ISR has been established for the subject-matter of claims 28, 29 and 51.
   Therefore, these claims cannot be subjected to IPE.
- Industrial applicability (Art.33(4) PCT)

Claims 32-38, 52, 61 and 62 relate to subject-matter considered by this Authority to be covered by the provisions of Rule 67.1(iv) PCT. Consequently, no opinion will be formulated with respect to the industrial applicability of the subject-matter of these claims (Art.34(4)(a)(i) PCT).

### Re Item V

Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Reference is made to the following documents:

D1: WO 99 47677 A (GENENTECH INC) 23 September 1999 (1999-09-23)

D2: WO 99 37671 A (NA SONGQING ;SONG HO YEONG (US); DOU

# INTERNATIONAL PRELIMINARY International application No. PCT/US99/30503 EXAMINATION REPORT - SEPARATE SHEET

SHENSHEN (US); LILLY CO ELI () 29 July 1999 (1999-07-29)

D3: EP-A-0 984 063 (LILLY CO ELI) 8 March 2000 (2000-03-08)

D4: WO 98 07832 A (UNIV HELSINKI LICENSING ; LUDWIG INST CANCER

RES (US)) 26 February 1998 (1998-02-26)

D5: WO 98 24811 A (ZYMOGENETICS INC) 11 June 1998 (1998-06-11)

### 2. Novelty (Art.33(2) PCT):

The prior art discloses various members of the VEGF family (see for example the description, page 1, line 21 to page 2, line 37, D4 which discloses VEGF-D, and D5 which discloses ZVEGF2). In contrast, the present application provides a novel member of this family, denominated VEGF-X and defined by its sequence according to claim 1, and/or by particular domains, such as the CUB domain according to claim 39-42 or the VEGF-like domain according to claim 39-42 or the VEGF-like

Accordingly, the subject-matter of the claims as far as subjected to IPE is novel.

### 3. Inventiveness (Art.33(3) PCT):

The novel member of the VEGF family is not rendered obvious by the cited prior art. Accordingly, the subject-matter of the claims as far as subjeted to IPE also comprises an inventive step.

#### Re Item VI

Certain documents cited
Certain published documents (Rule 70.10)

Application No
Patent No
Publication date
(day/monttr/year)
(day/monttr/year)
(day/monttr/year)
(day/monttr/year)

D1 WO9947677 23.09.99 10.03.99 17.03.98

02.11.98

D2 WO9937671 29.07.99 26.01.99

27.01.98 05.06.98

INTERNATIONAL PRELIMINARY

International application No. PCT/US99/30503

**EXAMINATION REPORT - SEPARATE SHEET** 

24.06.98

31.08.98

D3 EP-A-0 984063

08.03.00

16.07.99

31.08.98

### Re Item VII

Certain defects in the international application

Reference is made to claims 17, 18 and 65-67 which embrace human organism in their scope. This subject-matter is considered by the IPEA to be contrary to morality and are hence not allowable.

### Re Item VIII

Certain observations on the international application

The term "functional equivalent" embraces structurally undefined proteins. In so far the claims referring to this feature do not meet the requirements of Art.6 PCT (lack of clarity).

In addition, and since other VEGF molecules share properties of the VEGF-X peptide of the present application, the scope of the claims thereby also embraces proteins which are not part of the present invention (Art.6 PCT, lack of support).

It is also noted that the claims referring to this feature seem to embrace prior art VEGFs and would therefore also not meet the requirements of Art.33(2) PCT.

- 2. The feature "bioprecursor" has not been defined either, neither in the claims nor in the description. Therefore, the skilled person is not in the position to exemplify such a precursor (Art.5 and Art.6 PCT).
- Claims 32-38, 52, 61 and 62 comprise the treatment of various diseases. However, there is no proof for the applicability claimed to be found in the description for any of these various applications. In the absence of such a proof the subject-matter of these claims appears rather speculative and cannot be considered to meet the requirements of Art.5 PCT.